## onal Application No PCT/EP2004/009055

### INTERNATIONAL SEARCH REPORT

a. classification of subject matter 1PC 7 C12P21/00 C07K14/56 According to International Patent Classification (IPC) or to both national classification and IPC Minimum documentation searched (classification system followed by classification symbols) B. FIELDS SEARCHED C12P C07K IPC 7 Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) EPO-Internal, BIOSIS, EMBASE, WPI Data, PAJ C. DOCUMENTS CONSIDERED TO BE RELEVANT Relevant to claim No. Citation of document, with indication, where appropriate, of the relevant passages Category ° HART R A ET AL: "LARGE SCALE, IN SITU ISOLATION OF PERIPLASMIC IGF-I FROM E. Α COLI" BIO/TECHNOLOGY, NATURE PUBLISHING CO. NEW YORK, US, vol. 12, November 1994 (1994-11), pages 1113-1117, XP001153305 ISSN: 0733-222X the whole document -/--Patent family members are listed in annex. X Further documents are listed in the continuation of box C. "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the Special categories of cited documents : "A" document defining the general state of the art which is not considered to be of particular relevance invention "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone "E" earlier document but published on or after the international filing date "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art. document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed "&" document member of the same patent family Date of mailing of the international search report Date of the actual completion of the international search 05 -07- 2005 14 April 2005

Authorized officer

Schneider, P

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## INTERNATIONAL SEARCH REPORT

Interganal Application No
PC1/EP2004/009055

A FRENCH C ET AL: "DEVELOPMENT OF A SIMPLE METHOD FOR THE RECOVERY OF RECOMBINANT PROTEINS FROM THE ESCHERICHIA COLI PERIPLASM" ENZYME AND MICROBIAL TECHNOLOGY, STONEHAM, NA, US, vol. 19, no. 5, 1996, pages 332-338, XP0060886796 ISSN: 0141-0229 the whole document  A DALMORA S ET AL: "Analysis of recombinant human growth hormone directly in osmotic shock fluids" JOURNAL OF CHROMATOGRAPHY A, ELSEVIER SCIENCE, NL, vol. 782, no. 2, 10 October 1997 (1997-10-10), pages 199-210, XP004096275 ISSN: 0021-9673 the whole document  A WO 89/00201 A (THE UNIVERSITY OF DUNDEE) 12 January 1989 (1989-01-12) the whole document  A WO 89/00409 A (INSIGHT STRATEGY & MARKETING LTD; PELEG, YOAV; PANCER, ZEEV; INSIGHT B) 16 January 2003 (2003-01-16) the whole document			PCT/EP2004/009055
A FRENCH C ET AL: "DEVELOPMENT OF A SIMPLE METHOD FOR THE RECOVERY OF RECOMBINANT PROTEINS FROM THE ESCHERICHIA COLI PERIPLASM" ENZYME AND MICROBIAL TECHNOLOGY, STONEHAM, MA, US, vol. 19, no. 5, 1996, pages 332-338, XP000886796 ISSN: 0141-0229 the whole document  A DALMORA S ET AL: "Analysis of recombinant human growth hormone directly in osmotic shock fluids" JOURNAL OF CHROMATOGRAPHY A, ELSEVIER SCIENCE, NL, vol. 782, no. 2, 10 October 1997 (1997-10-10), pages 199-210, XP004096275 ISSN: 0021-9673 the whole document  A WO 89/00201 A (THE UNIVERSITY OF DUNDEE) 12 January 1989 (1989-01-12) the whole document  WO 03/004599 A (INSIGHT STRATEGY & MARKETING LTD; PELEG, YOAV; PANCER, ZEEV; INSIGHT B) 16 January 2003 (2003-01-16)	.(Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	Polographia alaim No.
METHOD FOR THE RECOVERY OF RECUMBINANT PROTEINS FROM THE ESCHERICHIA COLI PERIPLASM" ENZYME AND MICROBIAL TECHNOLOGY, STONEHAM, MA, US, vol. 19, no. 5, 1996, pages 332-338, XP000886796 ISSN: 0141-0229 the whole document  A DALMORA S ET AL: "Analysis of recombinant human growth hormone directly in osmotic shock fluids" JOURNAL OF CHROMATOGRAPHY A, ELSEVIER SCIENCE, NL, vol. 782, no. 2, 10 October 1997 (1997-10-10), pages 199-210, XP004096275 ISSN: 0021-9673 the whole document  A WO 89/00201 A (THE UNIVERSITY OF DUNDEE) 12 January 1989 (1989-01-12) the whole document  A WO 03/004599 A (INSIGHT STRATEGY & MARKETING LTD; PELEG, YOAV; PANCER, ZEEV; INSIGHT B) 16 January 2003 (2003-01-16)	ategory °	Citation of document, with indication, where appropriate, of the relevant passages	nelevant to claim No.
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MARKETING LTD; PELEG, YUAV; PANCER, ZEEV; INSIGHT B) 16 January 2003 (2003-01-16)	A	12 January 1989 (1989-01-12)	
	A	MARKETING LTD; PELEG, YUAV; PANCER, ZEEV; INSIGHT B) 16 January 2003 (2003-01-16)	





	at the Coffice chart
Box II Observations where certain claims were found unsearchable (Continu	uation of item 2 of first sheet)
This International Search Report has not been established in respect of certain claims under	Article 17(2)(a) for the following reasons:
1. Claims Nos.: because they relate to subject matter not required to be searched by this Authority,	namely:
Claims Nos.:     because they relate to parts of the International Application that do not comply with an extent that no meaningful International Search can be carried out, specifically:	the prescribed requirements to such
Claims Nos.:     because they are dependent claims and are not drafted in accordance with the sec	
Box III Observations where unity of invention is lacking (Continuation of It	em 3 of first sneet)
This International Searching Authority found multiple inventions in this international applica	ition, as follows:
see additional sheet	ì
As all required additional search fees were timely paid by the applicant, this interest searchable claims.	national Search Report covers all
2. As all searchable claims could be searched without effort justifying an additional of any additional fee.	fee, this Authority did not invite payment
As only some of the required additional search fees were timely paid by the appl covers only those claims for which fees were paid, specifically claims Nos.:	licant, this International Search Report
4. X No required additional search fees were timely paid by the applicant. Conseque restricted to the invention first mentioned in the claims; it is covered by claims N 1-12 (all completely), 14-19 (all partially)	ently, this International Search Report is los.:
Hemark of Frotest	s were accompanied by the applicant's protest.  the payment of additional search fees.

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## FURTHER INFORMATION CONTINUED FROM PCT/ISA/ 210

This International Searching Authority found multiple (groups of) inventions in this international application, as follows:

1. claims: 1-12 (all completely),14-19 (all partially)

A process to release a recombinant polypeptide of interest from the periplasm of the host cells by applying an osmotic shock directly on the host cells in the fermentation medium.

2. claims: 13,20-23 (all completely),14-19 (all partially)

A process for the purification of a recombinant interferon alpha 2 from a crude preparation of interferon alpha 2 by applying a specific sequence of several chromatographic steps as defined in claim 13 (i) to (v).

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### INTERNATIONAL SEARCH REPORT

Information on patent family members

Interr	nal Application No
PCT	EP2004/009055

Patent document cited in search report		Publication date		Patent family member(s)	Publication date
WO 8900201	A	12-01-1989	WO GB	8900201 A1 2206881 A	12-01-1989 18-01-1989
WO 03004599	Α	16-01-2003	WO	03004599 A2	16-01-2003